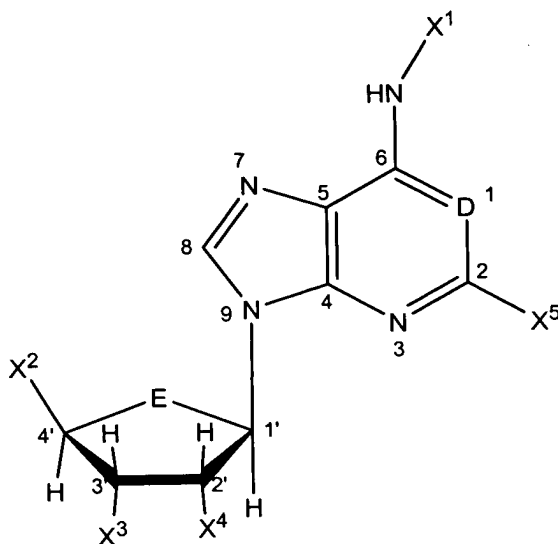


Amendments to the Claims:

1. (Currently amended) A product which is a compound of the formula:



wherein

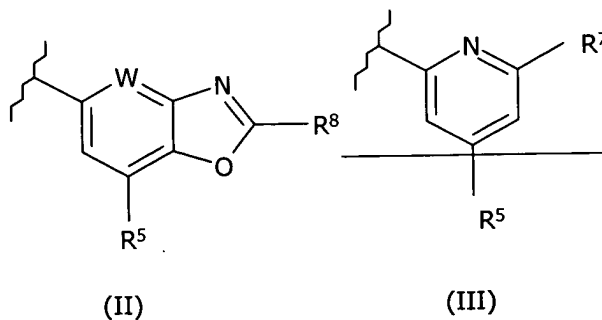
D is N or CH;

E is O, S or CH₂;

X¹ is a group of the formula -CR²⁰R²¹-CYCLE, where

R²⁰ and R²¹ are the same or different and H, F or CH₃;

CYCLE is of formula (II) ~~or formula (III)~~:



where:

R⁵ is iodine, bromine, methyl or trifluoromethyl;

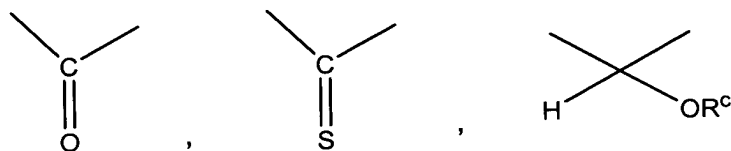
~~R⁷ is H, halogen, C₁-C₁₀-acyl, OR¹¹, CO₂R¹¹ or CONR¹¹ where R¹¹ is C₁-C₁₀-hydrocarbyl optionally containing one or more in-chain and/or in-ring O linkages;~~

R⁸ is -NR⁹R¹⁰ or -COR⁹, where R⁹ and R¹⁰ are each independently methyl or ethyl; and

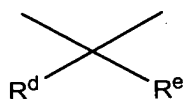
W is N or CH;

[.]X² is hydroxymethyl, (C₁-C₃)alkoxymethyl, (C₃-C₅)cycloalkoxy methyl, carboxy, (C₁-C₃)alkoxycarbonyl, (C₃-C₅)cycloalkoxy-carbonyl, 1,1-aminoiminomethyl, 1,1-(mono-N- or di-N,N-(C₁-C₄)alkylamino)iminomethyl, 1,1-(mono-N- or di-N,N-(C₃-C₅)cycloalkylamino)iminomethyl, carbamoyl, mono-N- or di-N,N-(C₁-C₄)alkylaminocarbonyl, mono-N- or di-N,N-(C₃-C₅)cycloalkyl-aminocarbonyl or N-(C₁-C₄)alkyl-N-(C₃-C₅)cycloalkylamino-carbonyl;

X³ and X⁴ are each independently hydrogen, alkyl, hydroxyalkyl, alkoxyalkyl, OR^a or NR^aR^b, where R^a and R^b are independently hydrogen, alkyl, aralkyl, carbamoyl, alkyl carbamoyl, dialkylcarbamoyl, acyl, alkoxycarbonyl, aralkoxycarbonyl, aryloxycarbonyl, or, when X³ and X⁴ are both OR^a, the two R^a groups together may form



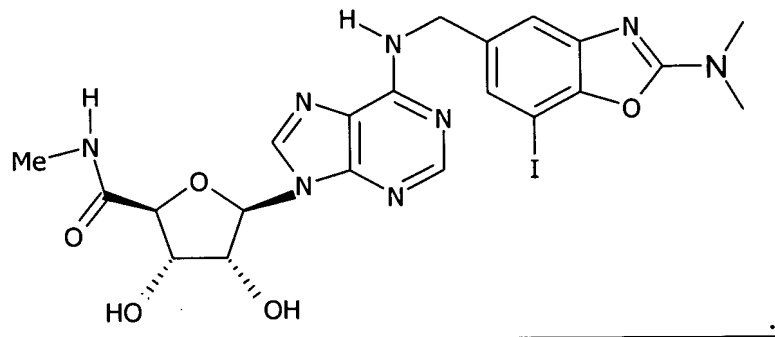
where R^c is hydrogen or alkyl,



where R^d and R^e are independently hydrogen, alkyl, or together with the carbon atom to which they are attached may form a 1,1-cycloalkyl group;

X⁵ is H, halogen, (C₁-C₁₀)alkyl, fluorinated (C₁-C₁₀) alkyl (e.g. trifluoromethyl), (C₁-C₁₀) alkoxyalkyl, (C₁-C₁₀)alkoxy, (C₁-C₁₀)alkylether, (C₁-C₁₀)thioalkoxy, (C₁-C₁₀)alkylthio, amino, (C₁-C₁₀)alkylamino, -COX⁶R²⁵ where X⁶ is O or NH and R²⁵ is (C₁-C₄)alkyl optionally terminally substituted by an aryl or a heteroaryl group and additionally or alternatively terminally substituted by hydroxy, (C₂-C₁₀)alkenyl, (C₂-C₁₀)alkynyl, or is (C₂-C₁₀)alkenyl, (C₂-C₁₀)alkynyl in either case terminally substituted by an aryl or heteroaryl group and, when having a terminal methylic carbon atom, optionally further terminally substituted by hydroxy,

or a pharmaceutically acceptable salt or prodrug thereof or a pharmaceutically acceptable salt of such a prodrug[[]], provided that the compound is not



2. (Original) A product of claim 1, wherein

D is N;

E is O;

X² is mono-N- or di-N,N(C₁-C₄)alkylaminocarbonyl, mono-N-

or di-, N-(C₃-C₅)cycloalkylaminocarbonyl or N-(C₁-C₄)alkyl-N- (C₃-C₅)cycloalkylaminocarbonyl;

X³ is OH or NH₂;

X⁴ is OH;

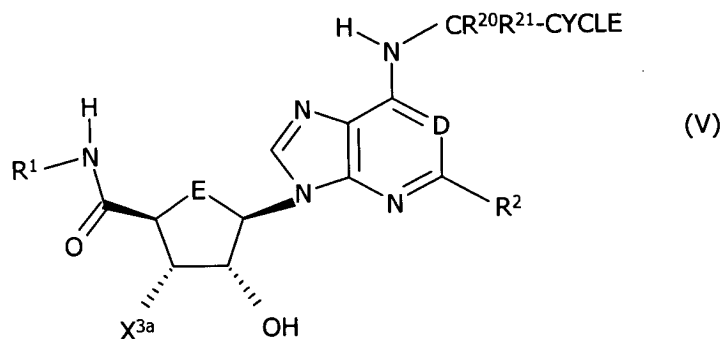
X⁵ is H, halogen, (C₁-C₁₀)alkyl, trifluoromethyl, (C₂-C₁₀)alkenyl, (C₂-C₁₀)alkynyl, or either of the latter two groups where terminally substituted as defined in claim 1.

3. (Currently amended) A product of claim 1 ~~or claim 2~~ wherein X⁵ is halogen.

4. (Currently amended) A product of claim ~~[[3]]~~ 2 wherein X⁵ is bromine or chlorine.

5. (Currently amended) A product of ~~any preceding~~ claim 1 wherein R²⁰ and R²¹ are both H.

6. (Currently amended) A product of claim 1 wherein the compound is of formula (V):



where:

-CR²⁰R²¹-CYCLE[[,]] and D and R² are as defined in claim 1;

R² is H, halogen, (C₁-C₁₀)alkyl, fluorinated (C₁-C₁₀) alkyl (e.g. trifluoromethyl), (C₁-C₁₀) alkoxyalkyl, (C₁-C₁₀)alkoxy, (C₁-C₁₀)alkylether, (C₁-C₁₀)thioalkoxy, (C₁-C₁₀)alkylthio, amino, (C₁-C₁₀)alkylamino, -COX⁶R²⁵ where X⁶ is O or NH and R²⁵ is (C₁-C₄)alkyl optionally terminally substituted by an aryl or a heteroaryl group and additionally or alternatively terminally substituted by hydroxy, (C₂-C₁₀)alkenyl, (C₂-C₁₀)alkynyl, or is (C₂-C₁₀)alkenyl, (C₂-C₁₀)alkynyl in either case terminally

substituted by an aryl or heteroaryl group and, when having a terminal methylic carbon atom, optionally further terminally substituted by hydroxy;

E is O, S or CH₂ (e.g. E is O and optionally D is N and R² is Cl or other halogen);

R¹ is C₁-C₄ alkyl; and

X^{3a} is -OH or -NH₂.

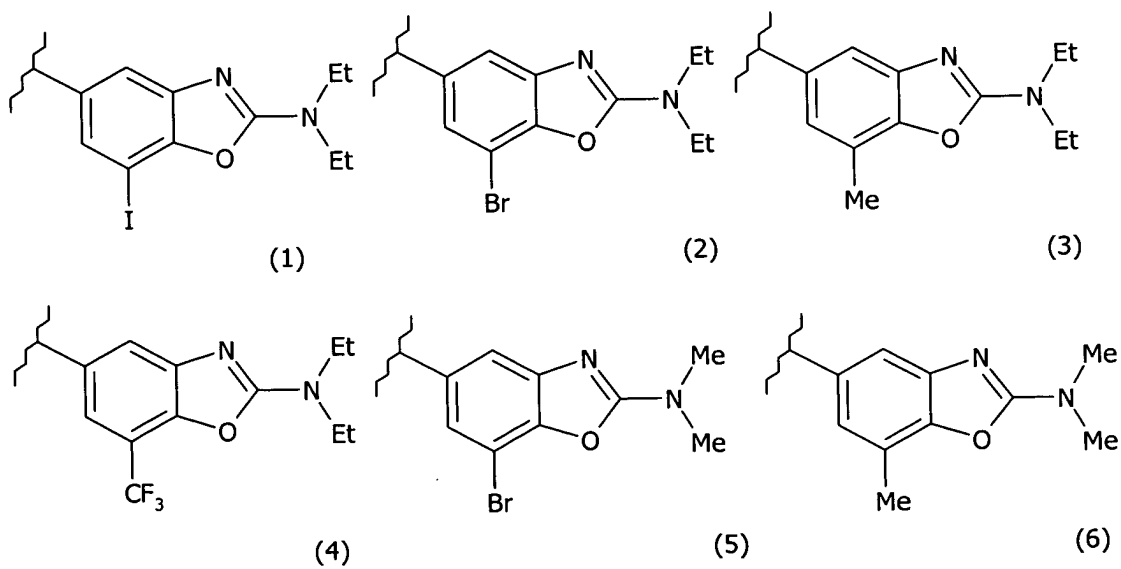
7. (Original) A product of claim 6 wherein E is O.

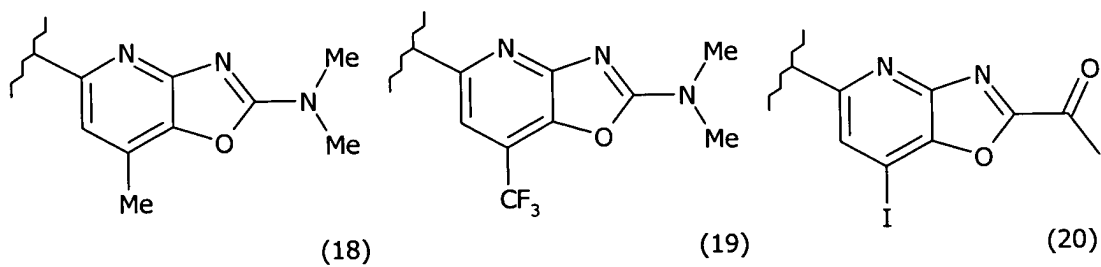
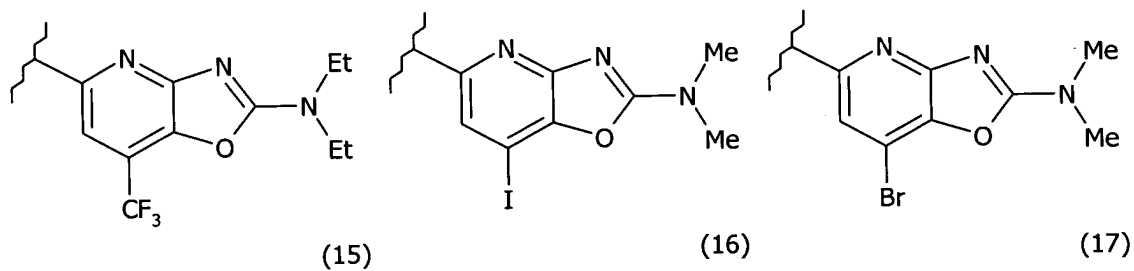
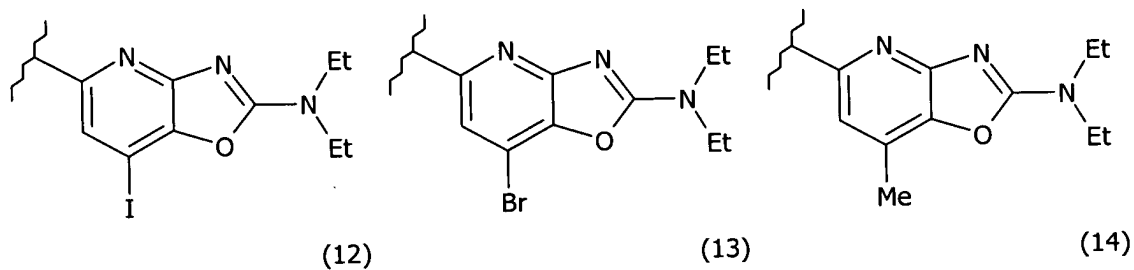
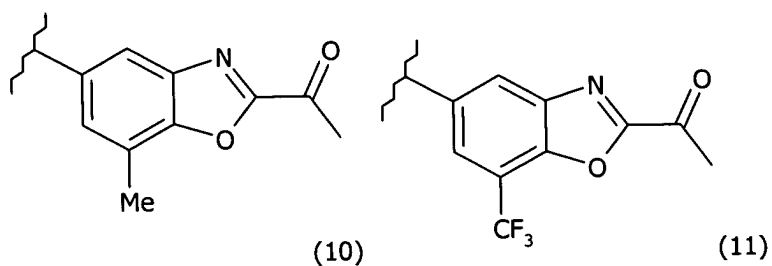
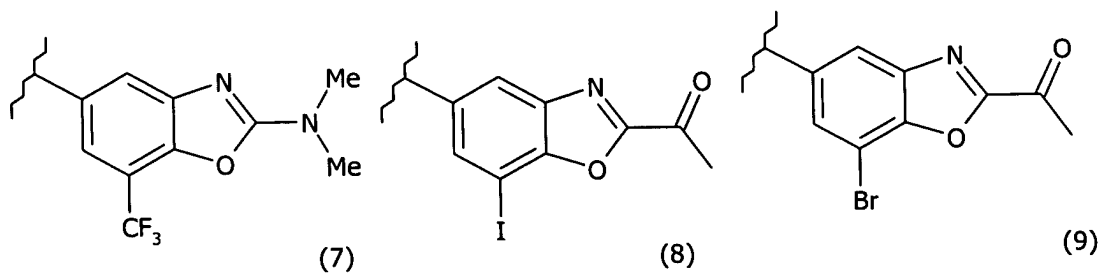
8. (Canceled)

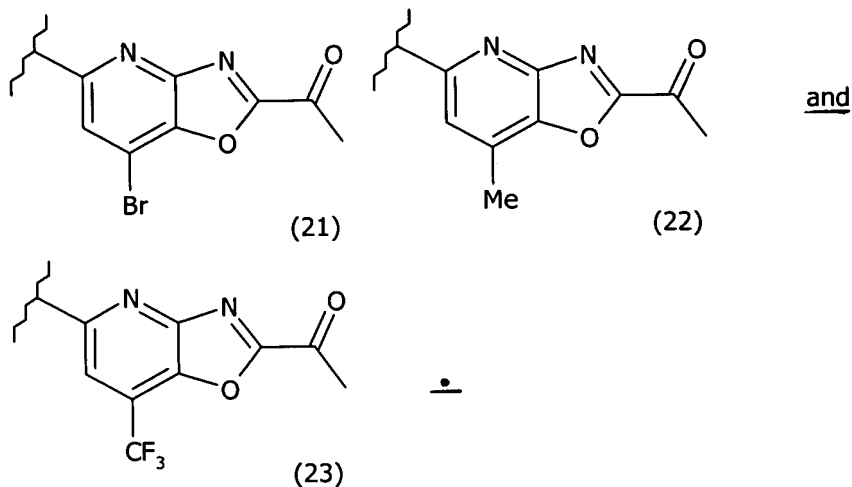
9. (Currently amended) A product of claim [[8]] 1 wherein W is N.

10. (Currently amended) A product of claim ~~8 or claim 9~~ 1 wherein R⁸ is dimethylamino or diethylamino.

11. (Currently amended) A product of claim [[8]] 1 wherein CYCLE is selected from the group consisting of the following moieties:

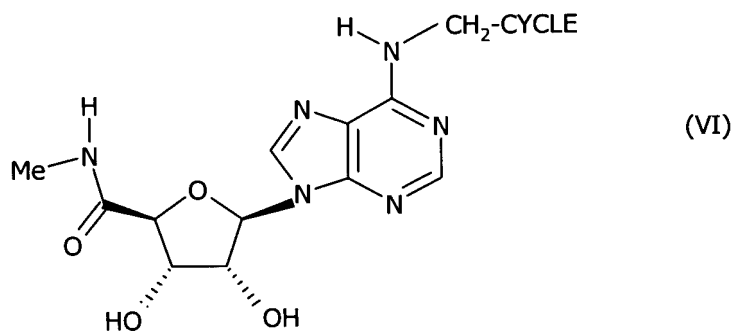






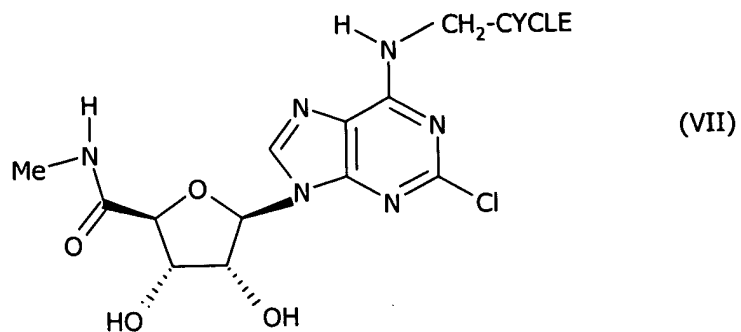
12. (Original) A product of claim 11 wherein CYCLE is of formula 1, 2, 3 or 4;
or of formula 12, 13, 14 or 15.

13. (Currently amended) A product of ~~any of claims 8, 11 and 12~~ claim 1
wherein the compound is of formula (VI):



where CYCLE is a group of formula (II).

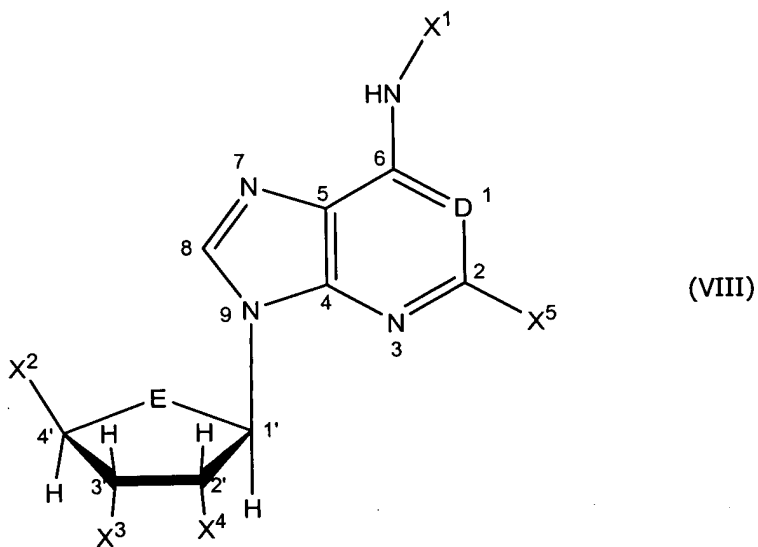
14. (Currently amended) A product of ~~any of claims 8, 11 and 12~~ claim 1
wherein the compound is of formula (VII):



where CYCLE is a group of formula (II).

15-19. (Canceled)

20. (Currently amended) A compound product which is a compound of formula (VIII):

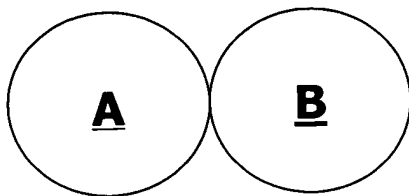


wherein

D is N or CH;

E is O, S or CH₂;

X^1 is of the formula $-CR^{20}R^{21}-CYCLE$ where R^{20} and R^{21} are the same or different and H, F or CH_3 ; and CYCLE is a bicyclic (fused) heteroaromatic ring of the formula



wherein

ring A is a 5- or 6- membered ring characterised by the following features (in which ring positions are numbered relative to the linkage to $-CR^{20}R^{21}-$):

- i. a carbon atom at the 1-position;
- ii. carbon atom as CH or a nitrogen atom at position 2;
- iii. it is 3, 4 fused to ring B;
- iv. the 5-position ring atom is substituted by a moiety R^5 which is H, halogen, or an organic moiety having from 1 to 6 plurally valent atoms in addition to monovalent atoms selected from hydrogen and halogen;
- v. if a 6-membered ring, it has at the 6-position a nitrogen, or $-CM-$ where M is H, CH_3 or F;

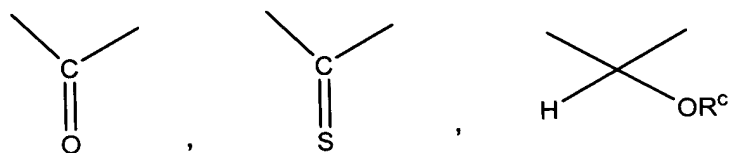
ring B is a 5 or 6 membered ring characterised by the following features:

- (a) an in-ring heteroatom including O, N or S joined to the 4-position of ring A;
- (b) said in-ring heteroatom is joined within the ring secondly to a carbon which is substituted by a moiety R^8 which is $-N(C_2H_5)_2$;
- (c) an in-ring atom joined to the 3-position of ring A which is N, O, S or C, said C being in the form of a CH or CO group;
- (d) in the case of a 6-membered ring, the remaining ring member is nitrogen or carbon in the form of CH;

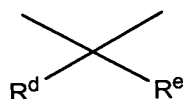
X^2 (the 4' substituent) is hydroxymethyl, (C_1-C_3) alkoxymethyl, (C_3-C_5) cycloalkoxy methyl, carboxy, (C_1-C_3) alkoxycarbonyl, (C_3-C_5) cycloalkoxycarbonyl, 1,1-aminoiminomethyl, 1,1-(mono-N- or di-N,N- (C_1-C_4) alkylamino)iminomethyl, 1,1-

(mono-N- or di-N,N-(C₃-C₅)cycloalkyl-amino)iminomethyl, carbamoyl, mono-N- or di-N,N-(C₁-C₄)alkylaminocarbonyl, mono-N- or di-N,N-(C₃-C₅)cycloalkylaminocarbonyl or N-(C₁-C₄)alkyl-N-(C₃-C₅)cycloalkylaminocarbonyl;

X³ and X⁴ are each independently hydrogen, alkyl, hydroxyalkyl, alkoxyalkyl, OR^aNR^aR^b, where R^a and R^b are independently hydrogen ~~(most preferably X³ and X⁴ are OH)~~, alkyl, aralkyl, carbamoyl, alkyl carbamoyl, dialkylcarbamoyl, acyl, alkoxy carbonyl, aralkoxy carbonyl, aryloxy carbonyl, or, when X³ and X⁴ are both OR^a, the two R^a groups together may form



where R^c is hydrogen or alkyl,



where R^d and R^e are independently hydrogen, alkyl, or together with the carbon atom to which they are attached may form a 1,1-cycloalkyl group;

X⁵ is H, halogen, (C₁-C₁₀)alkyl, fluorinated (C₁-C₁₀) alkyl (e.g. trifluoromethyl), (C₁-C₁₀) alkoxyalkyl, (C₁-C₁₀)alkoxy, (C₁-C₁₀)alkylether, (C₁-C₁₀)thioalkoxy, (C₁-C₁₀)alkylthio, amino, (C₁-C₁₀)alkylamino, -COX⁶R²⁵ where X⁶ is O or NH and R²⁵ is (C₁-C₄)alkyl optionally terminally substituted by an aryl or a heteroaryl group ~~{for example phenyl or a 5 or 6 membered heteroaryl group}~~ and additionally or alternatively terminally substituted by hydroxy, (C₂-C₁₀)alkenyl, (C₂-C₁₀)alkynyl, or is (C₂-C₁₀)alkenyl or (C₂-C₁₀)alkynyl in either case terminally substituted by an aryl or heteroaryl group ~~{for example phenyl or a 5 or 6 membered heteroaryl group}~~ and, when having a terminal methylic carbon atom, optionally further terminally substituted by

hydroxy[[.]], or a pharmaceutically acceptable salt or prodrug thereof, or a pharmaceutically acceptable salt of such a prodrug.

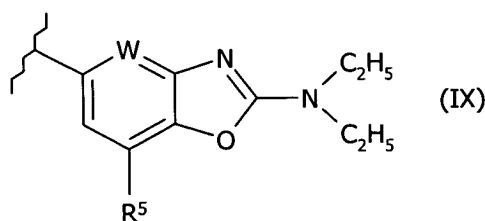
21. (Currently amended) A ~~compound~~ product of claim 20 wherein R⁵ has from 1 to 4 plurally valent atoms.

22. (Currently amended) A ~~compound~~ product of claim 21 wherein the plurally valent atoms are selected from carbon, oxygen, sulfur and nitrogen.

23. (Currently amended) A ~~compound~~ product of claim 22 wherein R⁵ is CH₃, CF₃, OH or NH₂.

24. (Currently amended) A ~~compound~~ product of claim 20 wherein R⁵ is H, I, Br or Cl.

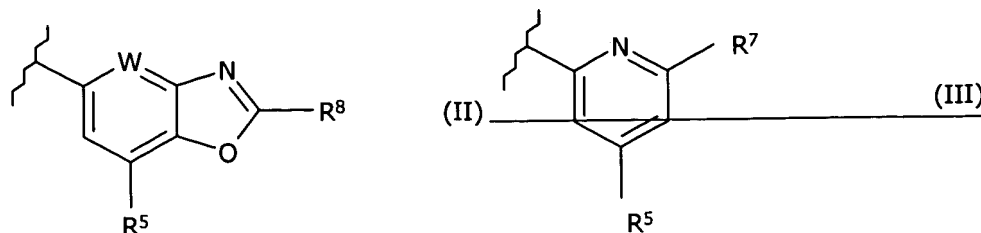
25. (Currently amended) A ~~compound~~ product of ~~any of~~ claim[[s]] 20 ~~to 24~~ wherein CYCLE is of formula (IX):



26. (Currently amended) A ~~compound~~ product of ~~any of~~ claim[[s]] 20 ~~to 25~~ wherein ~~where~~ R²⁰ and R²¹ are both hydrogen.

27. (Currently amended) An adenosine analogue-type A₃ receptor agonist having an N₆ nitrogen substituted by a group of the formula -CR²⁰R²¹-CYCLE where

R²⁰ and R²¹ are the same or different and H, F or CH₃; and CYCLE is of formula (II) or formula (III):



where:

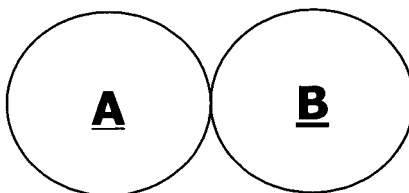
R⁵ is iodine, bromine, methyl or trifluoromethyl;

R⁷ is H, halogen, C₁-C₁₀-acyl, OR¹¹, CO₂R¹¹ or CONR¹¹ where R¹¹ is C₁-C₁₀ hydrocarbyl optionally containing one or more in chain and/or in ring O linkages;

R⁸ is -NR⁹R¹⁰ or -COR⁹, where R⁹ and R¹⁰ are each independently methyl or ethyl; and

W is N or CH.

28. (Original) An adenosine analogue-type A₃ receptor agonist having an N₆ nitrogen substituted by a group of the formula -CR²⁰R²¹-CYCLE where R²⁰ and R²¹ are the same or different and H, F or CH₃; and CYCLE is a bicyclic (fused) heteroaromatic ring of the formula



wherein

ring A is a 5- or 6- membered ring characterised by the following features (in which ring positions are numbered relative to the linkage to -CR²⁰R²¹-):

- i. a carbon atom at the 1-position;
 - ii. carbon atom as CH or a nitrogen atom at position 2;
 - iii. it is 3, 4 fused to ring B;
 - iv. the 5-position ring atom is substituted by a moiety R⁵ which is H, halogen or an organic moiety having from 1 to 6 plurally valent atoms in addition to monovalent atoms selected from hydrogen and halogen;
 - v. if a 6-membered ring, it has at the 6-position a nitrogen, or -CM- where M is H, CH₃ or F;
- ring B is a 5 or 6 membered ring characterised by the following features:
- (a) an in-ring heteroatom including O, N or S joined to the 4-position of ring A;
 - (b) said in-ring heteroatom is joined within the ring secondly to a carbon which is substituted by a moiety R⁸ which is -N(C₂H₅)₂;
 - (c) an in-ring atom joined to the 3-position of ring A which is N, O, S or C, said C being in the form of a CH or CO group;
 - (d) in the case of a 6-membered ring, the remaining ring member is nitrogen or carbon in the form of CH.

29. (Canceled)

30. (Currently amended) ~~A product of any one of claims 1 to 26 or an agonist of claim 27 or claim 28 for use in a method for selectively activating A₃ adenosine receptors in a mammal[[.]], comprising administering to the mammal an effective amount of a product of claim 1 or an agonist of claim 27.~~

31-32. (Canceled)

33. (Currently amended) ~~The use of a product of any one of claims 1 to 26 or an agonist of claim 27 or claim 28 for the manufacture of a medicament for use~~ A method for preconditioning the heart of a subject to protect it from ischaemic damage[[.]],

comprising administering to the subject an effective amount of a product of claim 1 or an agonist of claim 27.

34-35. (Canceled)

36. (Currently amended) A pharmaceutical composition comprising a product of ~~any one of claim[[s]] 1 to 26~~ or an agonist of claim 27 ~~or claim 28~~.

37. (Original) A pharmaceutical composition of claim 36 which is an intravenous formulation.

38. (Canceled)

39. (Currently amended) A method of stimulating adenosine A₃ receptors, comprising administering to a mammal in need of such treatment a therapeutically effective amount of a product of ~~any one of claim[[s]] 1 to 26~~ or an agonist of claim 27 ~~or claim 28~~.

40. (Currently amended) A method of reducing tissue or organ damage (~~e.g., substantially preventing tissue or organ damage, inducing tissue or organ protection~~) resulting from ischaemia or hypoxia, comprising administering to a mammal in need of such treatment a therapeutically effective amount of ~~an agent selected from a product of any one of claim[[s]] 1 to 26 and or an~~ agonist of claim 27 ~~or claim 28~~.

41. (New) The method of claim 39 wherein another cardiovascular drug is additionally administered to the mammal.